SEQUENCE LISTING <110> Nemerow, Glen R. Li, Erguang BIFUNCTIONAL MOLECULES AND VECTORS COMPLEXED THEREWITH FOR TARGETED <120> GENE DELIVERY <130> 22908-1228 <140> Herewith <141> 2001-07-10 <150> converted to a provisional from 09/613,017) <151> 2000-07-10 <160> 33 <170> FastSEQ for Windows Version 4.0 <210> 1 <211> 1516 <212> DNA <213> Mouse <220> <221> CDS <222> (28) ... (1395) <223> DAV-1 heavy chain, penton base monoclonal antibody <400> 1 cagacactga acacactgac tctaacc atg gga tgg agc tgg atc ttt ctc ttc $$\operatorname{Met}$$ Gly Trp Ser Trp Ile Phe Leu Phe 54 102 ctc ctg tca gga act gca ggc gtc cac tct gag gtc cag ctt cag cag

Leu 10	Leu	Ser	Ğİy	Thr	Ala 15	ĠĪу	Val	His	Ser	Glu 20	Val	Gln	Leu	Gln	Gln 25	
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						ttc Phe										198
						ctt Leu										246
						aac Asn 80										294
						aac Asn										342
						gtc Val										390





	ggc Gly															438
	tct Ser															486
	gtg Val 155															534
	gtg Val															582
	gct Ala															630
	ccc Pro															678
cac His	ccg Pro	gcc Ala 220	agc Ser	agc Ser	acc Thr	aag Lys	gtg Val 225	gac Asp	aag Lys	aaa Lys	att Ile	gtg Val 230	ccc Pro	agg Arg	gat Asp	726
	ggt Gly 235															765
	atc Ile															822
	aag Lys															870
	cag Gln															918
	caa Gln															966
	ctt Leu 315															1014
_	agg Arg	_		_	_	_			_							1062
	aaa Lys															1110
cct	ccc	aag	gag	cag	atg	gcc	aag	gat	aaa	gtc	agt	ctg	acc	tgc	atg	1158

Pro Pro Lys Glu Gln Met Ala Lys Asp Lys Val Ser Leu Thr Cys Met	
ata aca gac ttc ttc cct gaa gac att act gtg gag tgg cag tgg aat	1206
Ile Thr Asp Phe Phe Pro Glu Asp Ile Thr Val Glu Trp Gln Trp Asn 380 385 390	
ggg cag cca gcg gag aac tac aag aac act cag ccc atc atg gac aca Gly Gln Pro Ala Glu Asn Tyr Lys Asn Thr Gln Pro Ile Met Asp Thr 395 400 405	1254
gat ggc tct tac ttc gtc tac agc aag ctc aat gtg cag aag agc aac Asp Gly Ser Tyr Phe Val Tyr Ser Lys Leu Asn Val Gln Lys Ser Asn 410 415 420 425	1302
tgg gag gca gga aat act ttc atc tgc tct gtg tta cat gag ggc ctg Trp Glu Ala Gly Asn Thr Phe Ile Cys Ser Val Leu His Glu Gly Leu 430 435 440	1350
cac aac cac cat act gag aag agc ctc tcc cac tct cct ggt aaa His Asn His His Thr Glu Lys Ser Leu Ser His Ser Pro Gly Lys 445 450 455	1395
tgatcccagt gtccttggag ccctctggtc ctacaggact ctgtcaccta cctccacccc tccctgtata aataaagcac ctagcactgc cttgggaccc tgcaataaaa aaaaaaaaaa	1455 1515 1516
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Cys Thr Val Pro Glu Val Ser Ser Val Phe Ile Phe Pro Pro Lys Pro
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Phe Pro Ala Pro Ile Glu Lys Thr Ile Ser Lys Thr Lys Gly Arg Pro
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Lys Asn Thr Gln Pro Ile Met Asp Thr Asp Gly Ser Tyr Phe Val Tyr
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                                    410
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tgg gtt cca ggc tcc act ggt gac att gtg ctg acc caa tct cca gct
                                                                       99
Trp Val Pro Gly Ser Thr Gly Asp Ile Val Leu Thr Gln Ser Pro Ala
    15
tot ttg got gtg tot ota ggg cag agg god acc atc toc tgc aag god
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Ser Leu Āla Val Ser Leu Gly Gln Arg Āla Thr Ile Ser Cys Lys Āla
30
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55

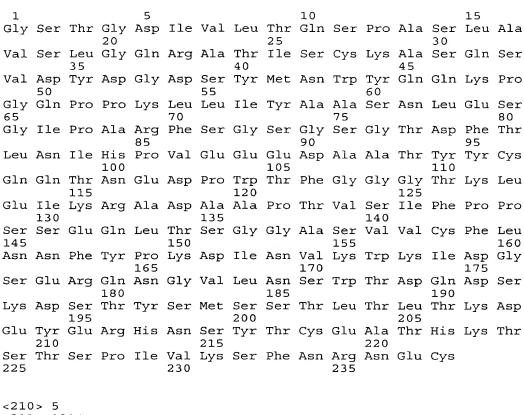
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tta gaa tct ggg atc cca gcc agg ttt agt ggc agt ggg tct ggg aca Leu Glu Ser Gly Ile Pro Ala Arg Phe Ser Gly Ser Gly Ser Gly Thr 80 85 90	291									
gac ttc acc ctc aac atc cat cct gtg gag gag gag gat gct gca acc Asp Phe Thr Leu Asn Ile His Pro Val Glu Glu Glu Asp Ala Ala Thr 95 100 105	339									
tat tac tgt cag caa act aat gag gat ccg tgg acg ttc ggt gga ggc Tyr Tyr Cys Gln Gln Thr Asn Glu Asp Pro Trp Thr Phe Gly Gly Gly 110 125	387									
acc aag ctg gaa atc aaa cgg gct gat gct gca cca act gta tcc atc Thr Lys Leu Glu Ile Lys Arg Ala Asp Ala Ala Pro Thr Val Ser Ile 130 135 140	435									
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tgt tagagacaaa ggtcctgaga cgccaccacc agctccccag ctccatccta Cys	776									
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<223> Portion of DAV-1 heavy chain used for fusion protein bifunctional antibody

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			aag Lys													288	1
			atg Met 100													336	÷
			gca Ala													384	:
			gca Ala													432	
			gct Ala													480	!
			tat Tyr													528	2
			agc Ser 180													576	
			ctg Leu	_	_				_			_				624	
			gtc Val													672	
			aaa Lys													720	
			cca Pro													768	
			ctc Leu 260													816	
			agc Ser													864	
			gag Glu													912	
			act Thr													960	
gac	tgg	ctc	aat	ggc	aag	gag	ttc	aaa	tgc	agg	gtc	aac	agt	gca	gct	1008	



Asp Trp Leu Asn Gly Lys Glu Phe Lys Cys Arg Val Asn Ser Ala Ala 325 330 335											
ttc cct gcc ccc atc gag aaa acc atc tcc aaa acc aaa ggc aga ccg Phe Pro Ala Pro Ile Glu Lys Thr Ile Ser Lys Thr Lys Gly Arg Pro 340 345 350	1056										
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aag gat aaa gtc agt ctg acc tgc atg ata aca gac ttc ttc cct gaa Lys Asp Lys Val Ser Leu Thr Cys Met Ile Thr Asp Phe Phe Pro Glu 370 375 380	1152										
gac att act gtg gag tgg cag tgg aat ggg cag cca gcg gag aac tac Asp Ile Thr Val Glu Trp Gln Trp Asn Gly Gln Pro Ala Glu Asn Tyr 385 390 395 400	1200										
aag aac act cag ccc atc atg gac aca gat ggc tct tac ttc gtc tac Lys Asn Thr Gln Pro Ile Met Asp Thr Asp Gly Ser Tyr Phe Val Tyr 405 410 415	1248										
agc aag ctc aat gtg cag aag agc aac tgg gag gca gga aat act ttc Ser Lys Leu Asn Val Gln Lys Ser Asn Trp Glu Ala Gly Asn Thr Phe 420 425 430	1296										
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Ser Leu Ser Ser Gly Val His Thr Phe Pro Ala Val Leu Gln Ser Asp
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                                                     190
Leu Tyr Thr Leu Ser Ser Ser Val Thr Val Pro Ser Ser Thr Trp Pro
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                            200
                                                 205
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Cys Thr Val Pro Glu Val Ser Ser Val Phe Ile Phe Pro Pro Lys Pro
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Lys Asp Val Leu Thr Ile Thr Leu Thr Pro Lys Val Thr Cys Val Val
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                                265
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Val Asp Ile Ser Lys Asp Asp Pro Glu Val Gln Phe Ser Trp Phe Val
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                            360
                                                 365
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                                             380
Asp Ile Thr Val Glu Trp Gln Trp Asn Gly Gln Pro Ala Glu Asn Tyr
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Lys Asn Thr Gln Pro Ile Met Asp Thr Asp Gly Ser Tyr Phe Val Tyr
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Ile Lys Ser Pro Cys Gln Arg Glu Thr Pro Glu Gly Ala Glu Ala Lys
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Pro Trp Tyr Glu Pro Ile Tyr Leu Gly Gly Val Phe Gln Leu Glu Lys
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Lys Pro Ala Lys Ser Ala
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<223> Stem Cell Factor (SCF, mature peptide)

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<211> 597

<212> PRT

<213> Artificial Sequence

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<223> Fusion protein with N-terminal portion of DAV-1 heavy chain and TNF alpha mature peptide

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                                         235
Cys Thr Val Pro Glu Val Ser Ser Val Phe Ile Phe Pro Pro Lys Pro
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                                     250
                                                         255
Lys Asp Val Leu Thr Ile Thr Leu Thr Pro Lys Val Thr Cys Val Val
                                265
            260
                                                     270
Val Asp Ile Ser Lys Asp Asp Pro Glu Val Gln Phe Ser Trp Phe Val
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                                                 285
Asp Asp Val Glu Val His Thr Ala Gln Thr Gln Pro Arg Glu Glu Gln
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                        295
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Phe Asn Ser Thr Phe Arg Ser Val Ser Glu Leu Pro Ile Met His Gln
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                                         315
Asp Trp Leu Asn Gly Lys Glu Phe Lys Cys Arg Val Asn Ser Ala Ala
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                                                     350
Lys Ala Pro Gln Val Tyr Thr Ile Pro Pro Pro Lys Glu Gln Met Ala
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                                             380
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                                         395
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Lys Asn Thr Gln Pro Ile Met Asp Thr Asp Gly Ser Tyr Phe Val Tyr
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                                     410
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                                                 445
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Gln Leu Gln Trp Leu Asn Arg Arg Ala Asn Ala Leu Leu Ala Asn Gly
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                                         475
Val Glu Leu Arg Asp Asn Gln Leu Val Val Pro Ser Glu Gly Leu Tyr
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Leu Ile Tyr Ser Gln Val Leu Phe Lys Gly Gln Gly Cys Pro Ser Thr
                                505
          500
                                                     51.0
His Val Leu Leu Thr His Thr Ile Ser Arg Ile Ala Val Ser Tyr Gln
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<213> Artificial Sequence

<220>

<223> Fusion protein with N-terminal portion of DAV-1 heavy chain and IGF-1 mature peptide

<400> 12

Met Gly Trp Ser Trp Ile Phe Leu Phe Leu Ser Gly Thr Ala Gly





1	•	_		5_					10					15	
Val	His	Ser	Glu 20	Val	GIn	Leu	GIn	G1n 25	Ser	GIY	Pro	Glu	Leu 30	Val	Lys
Pro	Gly	Ala 35	Ser	Val	Lys	Ile	Ser 40	Cys	Lys	Ala	Ser	Gly 45	Tyr	Thr	Phe
Thr	Asp 50	Tyr	Asn	Met	His	Trp 55	Val	Lys	Gln	Ser	His 60	Gly	Lys	Ser	Leu
Glu 65	Trp	Ile	Gly	Tyr	Ile 70	Tyr	Pro	Tyr	Lys	Gly 75	Gly	Thr	Gly	Tyr	Asn 80
			_	85	_				90			Ser		95	
		_	100			_		105			_	Ala	110		
		115					120					Gly 125			
	130				-	135					140	Tyr			
145					150					155		Leu			160
	_	_	-	165					170			Trp		175	-
			180	_				185				Leu	190		_
	-	195					200					Ser 205		-	
	210					215					220	Ser			
225					230					235		Lys			240
_				245					250			Pro		255	
			260					265		_		Thr	270		
	_	275		-	_	_	280					Ser 285	-		
_	290					295					300	Arg			
305					310					315		Ile			320
				325					330			Asn		335	
			340					345				Lys	350	-	
		355			_		360				_	Glu 365			
	370					375					380	Phe			Tyr
385					390		_		_	395		Tyr			400
				405					410			Gly		415	
			420			-		425	_			-	430		
	_	435					440	_				Leu 445	_	_	
	450		_			455				_	460	Arg	_		_
465					470					475		Arg			480
	_			485					490			Asp		495	Arg
neu	$\sigma_{\mathbf{L}}\mathbf{u}$	I'I'C'L	TAT	Cys	нта	PLO	ьeu	пув	PIO	нта	пуѕ	Ser	нта		

500 505

<210> 13 <211> 493

<212> PRT

<213> Artificial Sequence

<220>

<223> Fusion protein with N-terminal portion of DAV-1 heavy chain and EGF mature peptide

<400> 13 Met Gly Trp Ser Trp Ile Phe Leu Phe Leu Leu Ser Gly Thr Ala Gly 1.0 Val His Ser Glu Val Gln Leu Gln Gln Ser Gly Pro Glu Leu Val Lys 2.0 3.0 Pro Gly Ala Ser Val Lys Ile Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asp Tyr Asn Met His Trp Val Lys Gln Ser His Gly Lys Ser Leu Glu Trp Ile Gly Tyr Ile Tyr Pro Tyr Lys Gly Gly Thr Gly Tyr Asn Gln Lys Phe Lys Ser Lys Ala Thr Leu Thr Thr Asp Ser Ser Ser Asn Thr Ala Tyr Met Glu Leu Arg Ser Leu Thr Ser Asp Ala Ser Ala Val Tyr Tyr Cys Ala Arg Gly Ile Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ala Ala Lys Thr Thr Pro Pro Ser Val Tyr Pro Leu Ala Pro Gly Ser Ala Ala Gln Thr Asn Ser Met Val Thr Leu Gly Cys Leu 1.45 Val Lys Gly Tyr Phe Pro Glu Pro Val Thr Val Thr Trp Asn Ser Gly Ser Leu Ser Ser Gly Val His Thr Phe Pro Ala Val Leu Gln Ser Asp Leu Tyr Thr Leu Ser Ser Ser Val Thr Val Pro Ser Ser Thr Trp Pro Ser Glu Thr Val Thr Cys Asn Val Ala His Pro Ala Ser Ser Thr Lys Val Asp Lys Lys Ile Val Pro Arg Asp Cys Gly Cys Lys Pro Cys Ile Cys Thr Val Pro Glu Val Ser Ser Val Phe Ile Phe Pro Pro Lys Pro Lys Asp Val Leu Thr Ile Thr Leu Thr Pro Lys Val Thr Cys Val Val Val Asp Ile Ser Lys Asp Asp Pro Glu Val Gln Phe Ser Trp Phe Val Asp Asp Val Glu Val His Thr Ala Gln Thr Gln Pro Arg Glu Glu Gln Phe Asn Ser Thr Phe Arg Ser Val Ser Glu Leu Pro Ile Met His Gln Asp Trp Leu Asn Gly Lys Glu Phe Lys Cys Arg Val Asn Ser Ala Ala Phe Pro Ala Pro Ile Glu Lys Thr Ile Ser Lys Thr Lys Gly Arg Pro Lys Ala Pro Gln Val Tyr Thr Ile Pro Pro Pro Lys Glu Gln Met Ala Lys Asp Lys Val Ser Leu Thr Cys Met Ile Thr Asp Phe Phe Pro Glu Asp Ile Thr Val Glu Trp Gln Trp Asn Gly Gln Pro Ala Glu Asn Tyr





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Lys Asn Thr Gln Pro Ile Met Asp Thr Asp Gly Ser Tyr Phe Val Tyr
                405
                                    410
Ser Lys Leu Asn Val Gln Lys Ser Asn Trp Glu Ala Gly Asn Thr Phe
            420
                                425
                                                    430
Ile Cys Ser Val Leu His Glu Phe Asn Ser Asp Ser Glu Cys Pro Leu
                            440
                                                445
Ser His Asp Gly Tyr Cys Leu His Asp Gly Val Cys Met Tyr Ile Glu
                        455
                                            460
Ala Leu Asp Lys Tyr Ala Cys Asn Cys Val Val Gly Tyr Ile Gly Glu
                  470
                                      475
Arg Cys Gln Tyr Arg Asp Leu Lys Trp Trp Glu Leu Arg
                485
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<210> 14

<211> 613

<212> PRT

<213> Artificial Sequence

:220>

<400> 14

<223> Fusion protein with N-terminal portion of DAV-1 heavy chain and SCF mature peptide

Met Gly Trp Ser Trp Ile Phe Leu Phe Leu Leu Ser Gly Thr Ala Gly 10 Val His Ser Glu Val Gln Leu Gln Gln Ser Gly Pro Glu Leu Val Lys 25 Pro Gly Ala Ser Val Lys Ile Ser Cys Lys Ala Ser Gly Tyr Thr Phe 35 40 4.5 Thr Asp Tyr Asn Met His Trp Val Lys Gln Ser His Gly Lys Ser Leu 55 Glu Trp Ile Gly Tyr Ile Tyr Pro Tyr Lys Gly Gly Thr Gly Tyr Asn 70 75 Gln Lys Phe Lys Ser Lys Ala Thr Leu Thr Thr Asp Ser Ser Ser Asn 90 95 85 Thr Ala Tyr Met Glu Leu Arg Ser Leu Thr Ser Asp Ala Ser Ala Val 100 105 110 Tyr Tyr Cys Ala Arg Gly Ile Ala Tyr Trp Gly Gln Gly Thr Leu Val 115 120 125 Thr Val Ser Ala Ala Lys Thr Thr Pro Pro Ser Val Tyr Pro Leu Ala 135 140 Pro Gly Ser Ala Ala Gln Thr Asn Ser Met Val Thr Leu Gly Cys Leu 150 155 Val Lys Gly Tyr Phe Pro Glu Pro Val Thr Val Thr Trp Asn Ser Gly 165 170 Ser Leu Ser Ser Gly Val His Thr Phe Pro Ala Val Leu Gln Ser Asp 180 185 190 Leu Tyr Thr Leu Ser Ser Ser Val Thr Val Pro Ser Ser Thr Trp Pro 195 200 205 Ser Glu Thr Val Thr Cys Asn Val Ala His Pro Ala Ser Ser Thr Lys 215 220 Val Asp Lys Lys Ile Val Pro Arg Asp Cys Gly Cys Lys Pro Cys Ile 230 235 Cys Thr Val Pro Glu Val Ser Ser Val Phe Ile Phe Pro Pro Lys Pro 250 255 245 Lys Asp Val Leu Thr Ile Thr Leu Thr Pro Lys Val Thr Cys Val Val 260 265 Val Asp Ile Ser Lys Asp Asp Pro Glu Val Gln Phe Ser Trp Phe Val 275 280 285 Asp Asp Val Glu Val His Thr Ala Gln Thr Gln Pro Arg Glu Glu Gln 295 300 290 Phe Asn Ser Thr Phe Arg Ser Val Ser Glu Leu Pro Ile Met His Gln

23





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305
                    310
                                         315
Asp Trp Leu Asn Gly Lys Glu Phe Lys Cys Arg Val Asn Ser Ala Ala
                325
                                     330
Phe Pro Ala Pro Ile Glu Lys Thr Ile Ser Lys Thr Lys Gly Arg Pro
                                 345
                                                     350
Lys Ala Pro Gln Val Tyr Thr Ile Pro Pro Pro Lys Glu Gln Met Ala
        355
                             360
                                                 365
Lys Asp Lys Val Ser Leu Thr Cys Met Ile Thr Asp Phe Phe Pro Glu
    370
                        375
                                             380
Asp Ile Thr Val Glu Trp Gln Trp Asn Gly Gln Pro Ala Glu Asn Tyr
385
                    390
                                         395
Lys Asn Thr Gln Pro Ile Met Asp Thr Asp Gly Ser Tyr Phe Val Tyr
                                     410
                405
Ser Lys Leu Asn Val Gln Lys Ser Asn Trp Glu Ala Gly Asn Thr Phe
            420
                                 425
Ile Cys Ser Val Leu His Glu Phe Cys Arg Tyr Pro Ala Gln Trp Arg
                            440
        435
                                                 445
Pro Gln Gly Ile Cys Arg Asn Arg Val Thr Asn Asn Val Lys Asp Val
    450
                        455
                                            460
Thr Lys Leu Val Ala Asn Leu Pro Lys Asp Tyr Met Ile Thr Leu Lys
                                         475
                    470
Tyr Val Pro Gly Met Asp Val Leu Pro Ser His Cys Trp Ile Ser Glu
                485
                                     490
                                                         495
Met Val Val Gln Leu Ser Asp Ser Leu Thr Asp Leu Leu Asp Lys Phe
            500
                                 505
                                                     510
Ser Asn Ile Ser Glu Gly Leu Ser Asn Tyr Ser Ile Ile Asp Lys Leu
        515
                             520
                                                 525
Val Asn Ile Val Asp Asp Leu Val Glu Cys Val Lys Glu Asn Ser Ser
                        535
                                             540
Lys Asp Leu Lys Lys Ser Phe Lys Ser Pro Glu Pro Arg Leu Phe Thr
                    550
                                         555
Pro Glu Glu Phe Phe Arg Ile Phe Asn Arg Ser Ile Asp Ala Phe Lys
                565
                                     570
Asp Phe Val Val Ala Ser Glu Thr Ser Asp Cys Val Val Ser Ser Thr
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Leu Ser Pro Glu Lys Asp Ser Arg Val Ser Val Thr Lys Pro Phe Met
       595
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Leu Pro Pro Val Ala
    610
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     DAV-1 heavy chain.
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<212> DNA
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<400> 16

<220>

<223> PCR primer for amplification of CH1 region of

DAV-1 heavy chain.





cccagggtca tggagttag	19
<210> 17 <211> 20 <212> DNA <213> Artificial Sequence	
<220> <223> PCR primer for amplification of DAV-1 kappa chain CL-A.	
<400> 17 aagatggata cagttggtgc	20
<210> 18 <211> 20 <212> DNA <213> Artificial Sequence	
<220> <223> PCR primer for amplification of DAV-1 kappa chain CL-B.	
<400> 18 tgtcaagagc ttcaacagga	20
<210> 19 <211> 15 <212> PRT <213> Adenovirus	
<220> <221> PEPTIDE <222> (0)(0) <223> Peptide spanning integrin binding site on penton base.	
<400> 19 Met Asn Asp His Ala Ile Arg Gly Asp Thr Phe Ala Thr Arg Ala 1 5 10 15	
<210> 20 <211> 9 <212> PRT <213> Adenovirus	
<220> <221> PEPTIDE <222> (0)(0) <223> Epitope on penton base integrin binding site recognized by DAV-1	
<400> 20 Ile Arg Gly Asp Thr Phe Ala Thr Arg 1 5	
<210> 21 <211> 31 <212> DNA <213> Artificial Sequence	
<pre><220> <223> PCR sense primer for subcloning DAV-1 heavy chain for whole antil or Fab'2 constructs.</pre>	body





<400> ggtac	21 cgcca ccatgggatg gagctggatc t	31
<210><211><211><212><213>	24	
<220> <223>	PCR antisense primer for subcloning DAV-1 heavy chain for whole antibody construct.	
<400> gaatto	22 catgt aacacagagc agga	24
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<210><211><211><212><213>	28	
<220> <223>	PCR antisense primer for subcloning DAV-1 light chain for whole antibody or Fab'2 constructs.	
<400> tctaga	24 atgte tetaacaete atteetgt	28
<210><211><211><212><213>	24	
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<210> 27 <211> 26 <212> DNA <213> Artificial Sequence	
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<210> 28 <211> 26 <212> DNA <213> Artificial Sequence	
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<210> 30 <211> 96 <212> DNA <213> Artificial Sequence	
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<210><211><211><212><213>	26	
	PCR antisense primer for subcloning SCF into DAV-1/SCF fusion construct.	
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